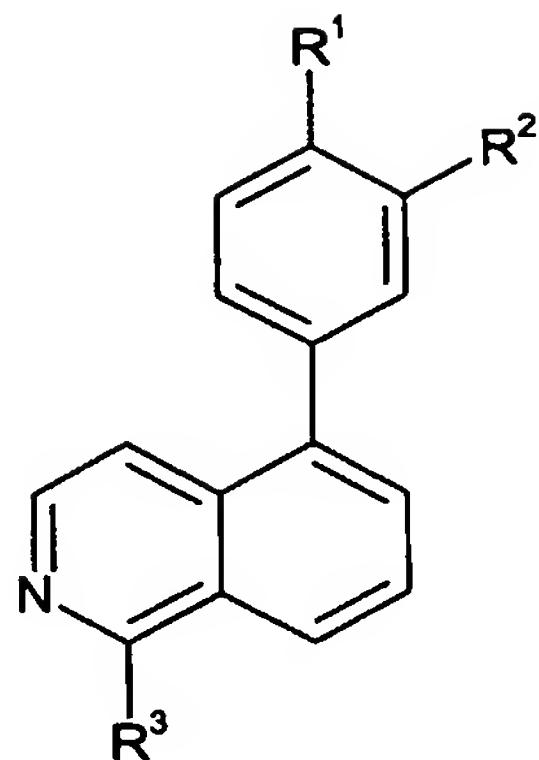


CLAIMS

1. A compound of Formula (I):



5 wherein:

One of R¹ and R² is H and the other represents –NHCONHR⁴

wherein R⁴ represents a phenyl or naphthyl group (which may be optionally substituted by one or more substituents independently selected from -C₁₋₆ alkyl, -C₁₋₆ haloalkyl, -CH₂CH₂CH₂-, halogen, C₁₋₆ alkoxy, C₁₋₆ haloalkoxy, OH, NO₂), C₃₋₇ cycloalkyl or R⁴ together with the NH to which it is bonded forms a morpholino group and

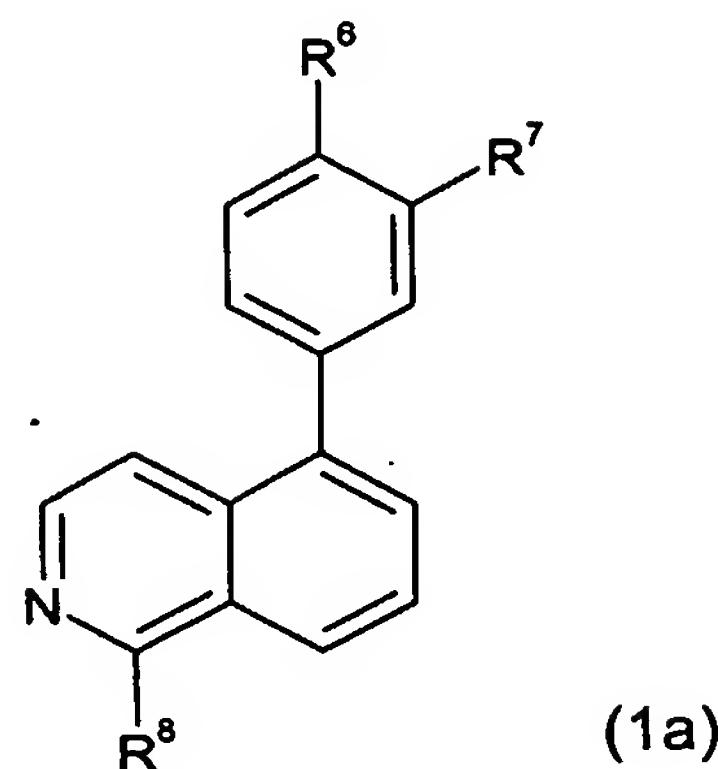
10 R³ is H or NHR⁵ wherein R⁵ is H, -quinolinyl or -isoquinolinyl, -(CONH)_p phenyl (wherein p is 0 or 1 and the phenyl is optionally substituted by one or more substituents independently selected from halogen, -C₁₋₆ alkyl, -C₁₋₆ haloalkyl, -morpholino, -SO₂NH₂, benzothiazole (substituted by methyl))

15 or a salt, solvate, or physiologically functional derivative thereof.

2. A compound according to claim 1 wherein R⁴ represents a phenyl group (which may be optionally substituted by one or more substituents selected from -C₁₋₆ haloalkyl, -CH₂CH₂CH₂-, halogen) or C₃₋₇ cycloalkyl.

20 3. A compound according to claims 1 – 2 wherein R³ is H or –NH R⁵ where in R⁵ is H, quinolinyl, -(CONH)p phenyl (wherein p is 0 or 1 and the phenyl is optionally substituted by one or more substituents independently selected from halogen, -C₁₋₆ haloalkyl –morpholino, -SO₂NH₂, benzothiazole, (substituted by methyl)).

4. A compound according to claims 1 – 3 of formula (1a)

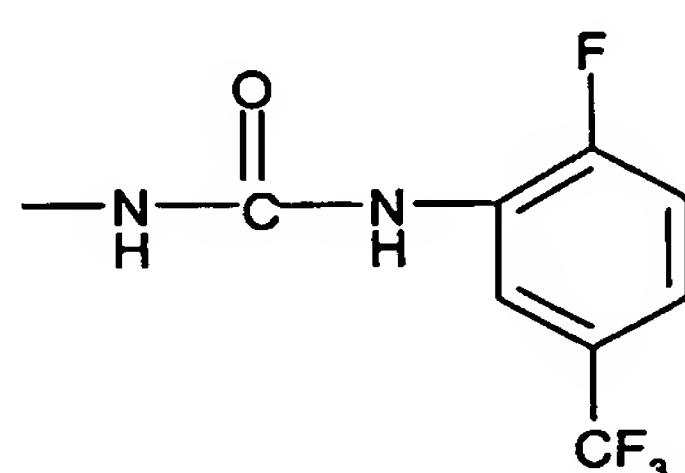


wherein one of R⁶ and R⁷ is H and the other represents -NHCONHR⁹;

5 R⁹ represents a phenyl group (which may be optionally substituted by one or more substituents independently selected from -C₁₋₆ haloalkyl, -CH₂CH₂CH₂-, halogen) or C₃₋₇ cycloalkyl;

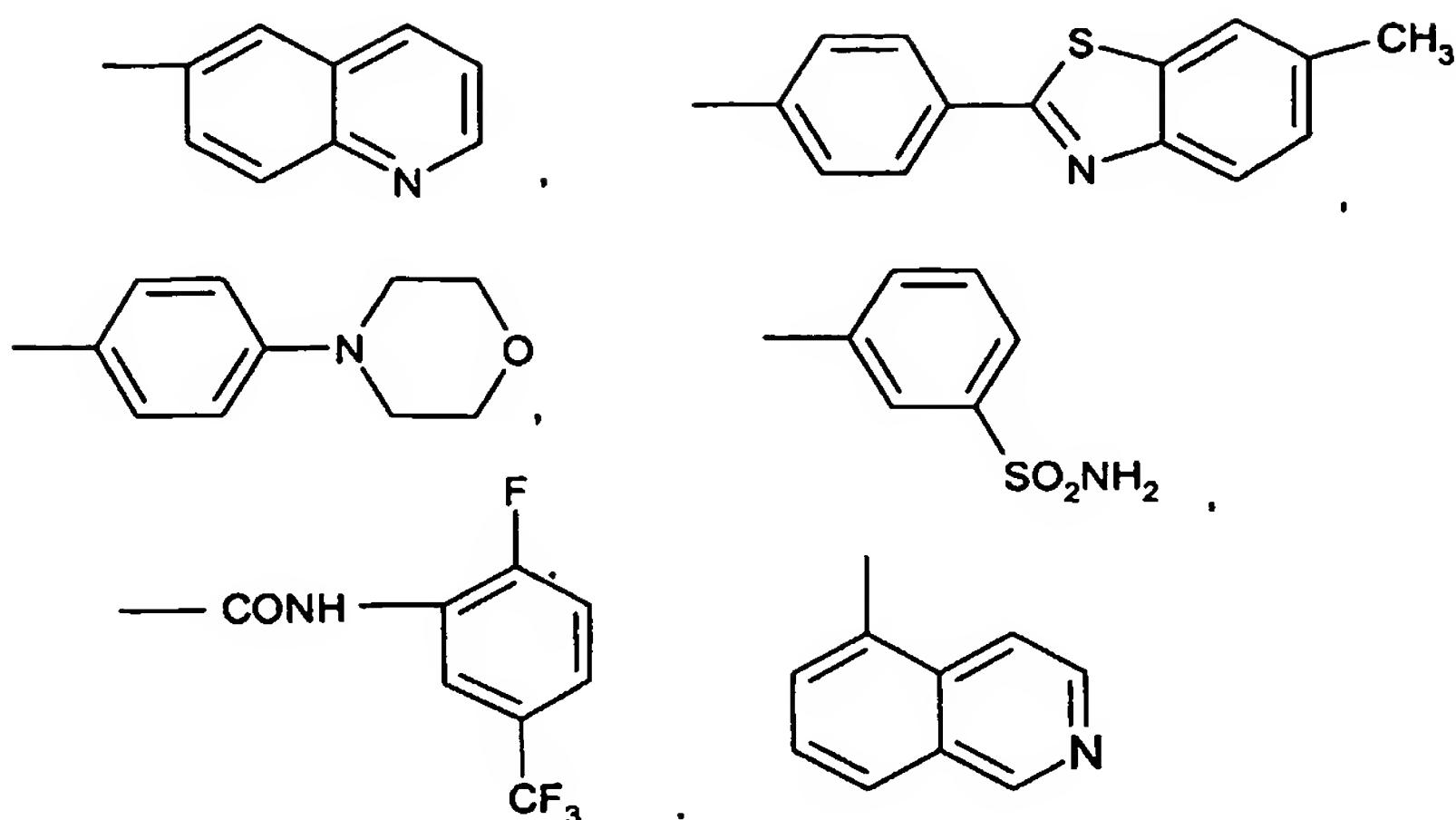
R⁸ is H or NHR¹⁰;

10 R¹⁰ is H quinolinyl, -(CONH)p phenyl (where p is 0 or 1 and the phenyl is optionally substituted by one or more substituents independently selected from halogen, -C₁₋₆ haloalkyl, -morpholino, -SO₂NH₂, benzothiazole (substituted by methyl)).

5. A compound according to claim 4 wherein NHCONHR⁹ represents

15

6. A compound according to claim 4 and 5 where in R¹⁰ is H,



7. A compound as claimed in claim 1 - 6, selected from the group consisting of:

1-(2-Fluoro-5-trifluoromethyl-phenyl)-3-(3-isoquinolin-5-ylphenyl)urea;

5 1-Cyclohexyl-3-(3-isoquinolin-5-ylphenyl)urea;

1-[3-(1-Amino-isoquinolin-5-yl)-phenyl]-3-(2-fluoro-5-trifluoromethyl-phenyl)-urea ;

1-(2-fluoro-5-trifluoromethyl-phenyl)-3-(5-{3-[3-(2-fluoro-5-trifluoromethyl-phenyl)-ureido]-phenyl}-isoquinolin-1-yl)-urea;

10 1-(2-Fluoro-5-trifluoromethyl-phenyl)-3-{3-[1-(quinolin-6-ylamino)-isoquinolin-5-yl]-phenyl}-urea;

1-(2-Fluoro-5-trifluoromethyl-phenyl)-3-(4-{1-[4-(6-methyl-benzothiazol-2-yl)-phenylamino]-isoquinolin-5-yl}-phenyl)-urea;

1-(2-Fluoro-5-trifluoromethyl-phenyl)-3-(3-{1-[4-(6-methyl-benzothiazol-2-yl)-phenylamino]-isoquinolin-5-yl}-phenyl)-urea;

15 1-(2-Fluoro-5-trifluoromethyl-phenyl)-3-(4-isoquinolin-5-ylphenyl)urea;

1-Indan-5-yl-3-(3-isoquinolin-5-yl-phenyl)-urea;

1-(2-Fluoro-5-trifluoromethyl-phenyl)-3-{3-[1-(4-morpholin-4-yl-phenylamino)-isoquinolin-5-yl]-phenyl}-urea;

3-{5-[3-(3-Cyclohexyl-ureido)-phenyl]-isoquinolin-1-ylamino}-benzenesulfonamide;

20

or a salt, solvate, or physiologically functional derivative thereof.

8. A pharmaceutical composition, comprising: a therapeutically effective amount

of a compound as claimed in any one of claims 1 - 7, or a salt, solvate, or a

25 physiologically functional derivative thereof and one or more of pharmaceutically acceptable carriers, diluents and excipients.

9. A pharmaceutical composition according to claim 8 further comprising an agent to inhibit growth factor receptor function
10. A compound as claimed in any of claims 1 - 7, or a salt, solvate, or a physiologically functional derivative thereof for use in therapy.
5
11. A method of treating a disorder in a mammal, said disorder being mediated by at least one of inappropriate TIE-2, Eph B4 and VEGFR-2 activity, comprising administering to said mammal a compound according to claims 1 - 7 or a salt, solvate or a physiologically functional derivative thereof.
10
12. The use of a compound according to claims 1 - 7, or a salt, solvate, or a physiologically functional derivative thereof in the manufacture of a medicament for use in the treatment of a disorder mediated by at least one of inappropriate TIE-2, EphB4 and VEGFR-2 activity.
15
13. A method of treating a disorder in a mammal, said disorder being mediated by at least one of inappropriate TIE-2, Eph B4 and VEGFR-2 activity, comprising: administering to said mammal (i) a compound according to claims 1 - 7, or a salt, solvate or physiologically functional derivative thereof and (ii) an agent to inhibit growth factor receptor function.
20
14. The use of a compound according to claims 1 - 7, or a salt, solvate or physiologically functional derivative thereof and an agent to inhibit growth factor receptor function in the manufacture of a medicament for the treatment of a disorder mediated by at least one of inappropriate TIE-2, EphB4 and VEGFR2 activity.
25
15. A method of treating a disorder in a mammal, said disorder being characterized by inappropriate angiogenesis, comprising administering to said mammal a compound according to claims 1 - 7, or a salt, solvate or physiologically functional derivative thereof.
30
16. The use of a compound according to claims 1 - 7 or a salt, solvate or physiologically functional derivative thereof in the manufacture of a medicament for the treatment of inappropriate angiogenesis.
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